

United States Patent and Trademark Office

UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER FOR PATENTS P.O. Box 1450 Alexandria, Virginia 22313-1450 www.uspto.gov

				•
APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/936,449	12/20/2001	Monica G. Marcu	213373	4132
45733 75	690 06/16/2006	EXAMINER		
LEYDIG, VOIT & MAYER, LTD. TWO PRUDENTIAL PLAZA, SUITE 4900 180 NORTH STETSON AVENUE			LE, EMILY M	
			ART UNIT	PAPER NUMBER
CHICAGO, IL	CHICAGO, IL 60601-6780			
			DATE MAILED: 06/16/2006	

Please find below and/or attached an Office communication concerning this application or proceeding.

	A martin attack	A 1! 4/. \			
	Application No.	Applicant(s)			
Office Action Summary	09/936,449	MARCU ET AL.			
Office Action Summary	Examiner	Art Unit			
	Emily Le	1648			
The MAILING DATE of this communication appears on the cover sheet with the correspondence address Period for Reply					
A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) FROM THE MAILING DATE OF THIS COMMUNICATION. - Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication. - If the period for reply specified above is less than thirty (30) days, a reply within the statutory minimum of thirty (30) days will be considered timely. - If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication. - Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).					
Status	•	•			
1) Responsive to communication(s) filed on 01/19	0/2006÷3/28/2006.				
	action is non-final.				
<u>'</u>	•	secution as to the merits is			
3) Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under <i>Ex parte Quayle</i> , 1935 C.D. 11, 453 O.G. 213.					
	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	0 0,0,2,0,			
Disposition of Claims	:				
4) ⊠ Claim(s) 1 and 3-17 is/are pending in the application. 4a) Of the above claim(s) is/are withdrawn from consideration. 5) □ Claim(s) is/are allowed. 6) ☒ Claim(s) 1 and 3-17 is/are rejected. 7) ☒ Claim(s) 1 and 3-17 is/are objected to. 8) □ Claim(s) are subject to restriction and/or election requirement.					
Application Papers					
9) The specification is objected to by the Examine	r				
10) ☐ The drawing(s) filed on is/are: a) ☐ accepted or b) ☐ objected to by the Examiner.					
Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).					
Replacement drawing sheet(s) including the correction is required if the drawing(s) is objected to. See 37 CFR 1.121(d).					
11) The oath or declaration is objected to by the Examiner. Note the attached Office Action or form PTO-152.					
Priority under 35 U.S.C. § 119					
 12) Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f). a) All b) Some * c) None of: 1. Certified copies of the priority documents have been received. 2. Certified copies of the priority documents have been received in Application No. 3. Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)). * See the attached detailed Office action for a list of the certified copies not received. 					
Attachment(s)	:				
1) Notice of References Cited (PTO-892)	4) Interview Summary	(PTO-413)			
2) Notice of Draftsperson's Patent Drawing Review (PTO-948)	Paper No(s)/Mail Da	ite			
3) Information Disclosure Statement(s) (PTO-1449 or PTO/SB/08) Paper No(s)/Mail Date	5)	atent Application (PTO-152)			

Art Unit: 1648

DETAILED ACTION

Continued Examination Under 37 CFR 1.114

1. A request for continued examination under 37 CFR 1.114, including the fee set forth in 37 CFR 1.17(e), was filed in this application after final rejection. Since this application is eligible for continued examination under 37 CFR 1.114, and the fee set forth in 37 CFR 1.17(e) has been timely paid, the finality of the previous Office action has been withdrawn pursuant to 37 CFR 1.114. Applicant's submissions filed on 01/19/2006 and 03/28/2006 has been entered.

Status of Claim(s)

2. Claims 2 and 18-23 are cancelled. Claims 1 and 3-17 are pending and under examination.

Claim Objections

3. Claims 1 and 3-17 are objected to because of the following informalities: the term coumarin is misspelled at line 6 of claim 1. Currently, the claim refers to coumarin derivatives as "coumain derivate". Appropriate correction is required.

Claim Rejections - 35 USC § 112

- 4. The following is a quotation of the second paragraph of 35 U.S.C. 112:
 The specification shall conclude with one or more claims particularly pointing out and distinctly claiming the subject matter which the applicant regards as his invention.
- 5. Claims 1 and 3-17 are rejected under 35 U.S.C. 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant regards as the invention.

র্

Application/Control Number: 09/936,449

Art Unit: 1648

The claims require the method to consist essentially of contacting a chaperone protein with coumarin or coumarin derivative.

Page 3

MPEP § 2111.02 [R-3] provides, the transitional term "consisting essentially of" limits the scope of the claim to the specified step, and those that do not materially affect the basic and novel characteristics of the claimed invention.

In the instant, the Office recognizes that the claimed method is limited to the specified step, contacting a chaperone protein with coumarin; however, it is not readily apparent from the claims and the specification what or which unspecified steps should be excluded to preserve the basic and novel characteristics of the claimed invention. This is particularly true because neither the specification nor the claims readily provide a clear teaching or clearly indicate the basic and novel characteristics that should be preserved with the claimed invention. Thus, in the absence of a clear indication from the specification and the claims of what the basic and novel characteristics actually are, the claims are rendered indefinitely because the metes and bounds of the claims are not readily be determined.

- 6. The following is a quotation of the first paragraph of 35 U.S.C. 112:
 - The specification shall contain a written description of the invention, and of the manner and process of making and using it, in such full, clear, concise, and exact terms as to enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the same and shall set forth the best mode contemplated by the inventor of carrying out his invention.
- 7. Claims 1 and 3-17 are rejected under 35 U.S.C. 112, first paragraph, as failing to comply with the written description requirement. The claim(s) contains subject matter which was not described in the specification in such a way as to reasonably convey to

Application/Control Number: 09/936,449

Art Unit: 1648

one skilled in the relevant art that the inventor(s), at the time the application was filed, had possession of the claimed invention. This is a new matter rejection.

Claim 1 has been amended to require the amount to be administered to "about 100 mg/kg", and at an interval of "at least once per day for about 5 days".

To show support for the cited recitations, Applicant cited lines 1-24 on page 16.

Lines 1-24 on page 16 provide the following: "...normal C57B16 mice received intraperitoneal injections (100 mg/kg) of novobiocin (a coumarin antibiotic) at 12 hour interval for 5 days".

Accordingly, the limitations 100 mg/kg and 12 hour interval for 5 days is fully supported by the passage cited by Applicant, and as recaptured above by the Office. However, the cited passage does not express, implicit or inherent, an administration dosage amount of "about 100 mg/kg", and regimen of "at least once per day for about 5 days", as currently recited in the claims. In the instant, a difference in scope exists between the dosage amount and frequency of the treatment regimen disclosed in the original specification and those recited in the current claims. Thus, in view of this difference in scope, the claims are rejected under 35 U.S.C 112, 1st paragraph for the absence of adequate written description for the cited recitations.

Additionally, claims 14-17 are also rejected under 35 U.S.C. 112, first paragraph, as failing to comply with the written description requirement for the presence of new matter. In the instant, the Office notes that claims 14-17 are not part of the originally filed specification. The originally filed specification contains only 13 claims, claims 1-13. In the instant, the Office has reviewed the disclosure of the international application,

Art Unit: 1648

from which the instant application claims 371 benefit to, and the specification of the instant application; however, neither implicit nor inherent support can be found with either disclosures for the language recited in claims 14-17. Thus, in the absence of expressed support, either implicit or inherent, for the language recited in claims 14-17, the claims are rejected under 35 U.S.C. 112, first paragraph, as failing to comply with the written description requirement for the presence of new matter.

Regarding the above-stated issue, the Office directs Applicant's attention to MPEP § 602 [R-3] (V), which provides the following: For applications filed prior to September 21, 2004, a preliminary amendment that is present on the filing date of an application may be considered a part of the original disclosure if it is referred to in a first filed oath or declaration in compliance with 37 CFR 1.63. If the preliminary amendment was not referred to in the oath or declaration, applicant will be required to submit a supplemental oath or declaration under 37 CFR 1.67 referring to both the application and the preliminary amendment filed with the original application. A surcharge under 37 CFR 1.16(f) will also be required unless it has been previously paid.

Claim Rejections - 35 USC § 102

8. The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless -

(b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.

9. Claims 1, 3 and 5-15 are rejected under 35 U.S.C. 102(b) as being anticipated by Eder et al., 1 as evidenced by Marcu et al. 2

The claims are directed to a method of inhibiting the binding of heat shock protein (Hsp) 90 with its client protein or client polypeptide in a mammal, wherein the method comprises contacting Hsp90 with coumarin or a coumarin derivative, whereby about 100 mg/kg of coumarin or coumarin derivative is administered to the mammal at least once per day for about 5 days.

Claim 3, which depends on claim 1, requires the coumarin or coumarin derivative to be a coumarin antibiotic. Claim 5, which depends on claim 3, defines the coumarin antibiotic as novobiocin. Claim 6, which depends on claim 1, also limits the coumarin or coumarin derivative to novobiocin. Claim 7, which depends on claim 6, requires novobiocin to bind to the carboxyl-terminal region of Hsp90. Claim 8, which depends on claim 1, defines the client protein or client polypeptide as tyrosine or serine/threonine kinase. Claim 9, which depends on claim 8, limits the tyrosine kinase to p185^{erB2} or p60^{v-src}. Claim 10, which depends on claim 8, limits the serine/threonine kinase to Raf-1. Claim 11, which depends on claim 1, defines the client protein or client polypeptide as mutated p53 protein. Claim 12, which depends on claim 1, expresses that the client protein or client polypeptide to be inactive subsequent to binding of Hsp90 to the coumarin or coumarin derivative. Claim 13, which depends on claim 12, specifies that the client protein or client polypeptide be degraded. Claim 14, which depends on claim

¹ Eder et al. Effect of novobiocin on the antitumor activity and tumor cell and bone marrow survivals of three alkylating agents. Cancer Research, 1989, Vol. 49, Issue 3, 595-598.

1, requires that Hsp90 be in a cell, and cellular proliferation is inhibited. Claim 15, which depends on claim 14, specifies that cellular proliferation is cancer.

For the purposes of searching for and applying prior art under 35 U.S.C. 102 and 103, the phrase "consisting essentially of" will be construed as equivalent to "comprising" since a clear indication of what the basic and novel characteristics actually are cannot be ascertained from the specification or the claims. See MPEP 2111.03 [R-3], 4th paragraph; and PPG, 156 F.3d at 1355, 48 USPQ2d at 1355, see also AK Steel Corp. v. Sollac, 344 F.3d 1234, 1240-41, 68 USPQ2d 1280, 1283-84 (Fed. Cir. 2003).

In the instant, Eder et al. teaches a method comprising the administration coumarin or coumarin derivative to a mammal. The coumarin derivative that Eder et al. administered to the mammal is novobiocin, a coumarin antibiotic. [Abstract] The amount of novobiocin that Eder et al. administered to the mammal includes 50 mg/kg and 100mg/kg to the mammals. [Tumor Growth Delay Experiments segment under the Materials and Methods section disclosed on page 595.] Eder et al. teaches the administration novobiocin once daily on days 6 through 15 post tumor implantation. In the instant, the frequency in which Eder et al. administered novobiocin is once daily for 10 days; wherein the 6th day post tumor implantation serves as the first day in which the first dose of novobiocin is administered and the 15th day post tumor implantation serves as the 10th day in which the last dose of novobiocin is administered to the mammal. Eder et al. notes that the administration of novobiocin inhibits cellular proliferation, particularly cancer.

² Marcu et al. The heat shock protein 90 antagonist novobiocin interacts with previously unrecognized ATP-binding domain in the carboxyl terminus of the chaperone. The Journal of Biological Chemistry,

In summary, Eder et al. teaches a method comprising the administration coumarin or coumarin derivative to a mammal. The amount in which novobiocin was administered to the mammal by Eder et al. is about 100 mg/kg. The frequency in which novobiocin is administered to the mammal by Eder et al. is at least once per day for about 5 days. In the instant, while the Office does note that the claims specifically recite "about 100 mg/kg" and "about 5 days"; however, in the absence of any guidance from specification for the extent of coverage the term "about" provides, the Office regards the 50 mg/kg and 100mg/kg, and the 10 day treatment period as being encompassed by recitations "about 100 mg/kg" and "about 5 days", respectively. Therefore, Eder et al. teaches a method comprising the administration coumarin or coumarin derivative to a mammal, wherein about 100 mg/kg of coumarin or coumarin derivative is administered to the mammal at least once per day for about 5 days, and whereby the coumarin or coumarin derivative is novobiocin.

Additionally, the Office notes that Eder et al. does not acknowledge that the administration of novobiocin inhibits the binding of Hsp90 with its client protein or client polypeptide. However, MPEP § 2112 [R-3] (I) provides the following:

"[T]he discovery of a previously unappreciated property of a prior art composition, or of a scientific explanation for the prior art's functioning, does not render the old composition patentably new to the discoverer." Atlas Powder Co. v. Ireco Inc., 190 F.3d 1342, 1347, 51 USPQ2d 1943, 1947 (Fed. Cir. 1999). Thus the claiming of a new use, new function or unknown property which is inherently present in the prior art does not necessarily make the claim patentable.

Thus, in accordance with the guidance provided by MPEP § 2112 [R-3] (I), Eder et al. does not need to appreciate or acknowledge that novobiocin inhibits the binding of

Hsp90 with its client protein or client polypeptide to render the claimed invention unpatentable. In the instant, while the Office does appreciate the scientific explanation offered by Applicant, however, Applicant's discovery or appreciation of the mechanism of action in which novobiocin function in a mammal is not sufficient to render Applicant's discovery patentable. In all, Applicant's appreciation or discovery of an unknown property that is inherently present in the prior art does not make Applicant's claimed invention patentable.

Furthermore, it is clearly evident from the art that novobiocin inhibits the binding of Hsp90 with its client protein or client polypeptide. See the teachings of Marcu et al. Marcu et al. teaches that novobiocin antagonizes Hsp90 in vitro and in vivo by binding to the carboxyl terminus of Hsp90. [First sentence under the Discussion section disclosed on page 37185, and the title of the article.] Marcu et al. further teaches that the antagonizing activity of novobiocin against Hsp90 leads to the rapid destabilization of various Hsp90 client proteins, including Raf-1, p60^{v-src}, and p185^{erbB2} --all of which are tyrosine and serine/threonin kinases, and mutated p53. [First full paragraph disclosed on the right column of page 371811 Thus, Marcu et al. clearly evidenced that the administration of novobiocin, as Eder et al. teaches, inherently lead to the inhibition of binding of Hsp90 with client protein or client polypeptides. Additionally, Marcu et al. clearly evidenced that novobiocin inherently i) binds to the carboxyl-terminal region of Hsp90; ii) inhibits the binding of Hsp90 with its client proteins (Raf-1, p185^{erbB2} and p60^{v-} src-- all of which are tyrosine and serine/threonin kinases, and mutated p53); and iii) inactivates the client proteins since the chaperone protein (Hsp90) is no longer available

Art Unit: 1648

to mediate the function of these client proteins; which in essence is the equivalent of degrading these proteins. Hence, in view of the evidence provided by Marcu et al., Eder et al. does teach a method of inhibiting the binding of Hsp90 with its client protein or client polypeptide in a mammal, wherein the method comprises contacting Hsp90 with novobiocin by administering about 100 mg/kg of novobiocin, for at least once per day for about 5 days to the mammal. Therefore, Eder et al. anticipates the claimed invention.

Claim Rejections - 35 USC § 103

- 10. The following is a quotation of 35 U.S.C. 103(a) which forms the basis for all obviousness rejections set forth in this Office action:
 - (a) A patent may not be obtained though the invention is not identically disclosed or described as set forth in section 102 of this title, if the differences between the subject matter sought to be patented and the prior art are such that the subject matter as a whole would have been obvious at the time the invention was made to a person having ordinary skill in the art to which said subject matter pertains. Patentability shall not be negatived by the manner in which the invention was made.
- 11. Claims 1 and 16-17 are rejected under 35 U.S.C. 103(a) as being unpatentable over Civitico et al.,³ as evidenced by Hu et al.⁴

The claims are directed to a method of inhibiting the binding of heat shock protein (Hsp) 90 with its client protein or client polypeptide in a mammal, wherein the method comprises contacting Hsp90 with coumarin or a coumarin derivative, whereby about 100 mg/kg of coumarin or coumarin derivative is administered to the mammal at least once per day for about 5 days. Claim 16, which depends on claim 1, requires the client protein to be hepatitis B virus reverse transcriptase. Claim 17, which depends on

³ Civitico et al. Antiviral strategies in chronic hepatitis B virus infection: II. Inhibition of duck hepatitis B virus in vitro using conventional antiviral agents and supercoiled-DNA active compounds. Journal of Medical virology, 1990, Vol. 31, 90-97.

claim 16, requires that the inhibition of binding of Hps90 with be hepatitis B virus reverse transcriptase results in the inhibition of hepatitis B viral replication.

In the instant, Civitico et al. teaches that novobiocin inhibits the replication of duck hepatitis B virus. Civitico et al. does not teach the administration of novobiocin to a mammal. However, Civitico et al. does suggest the use of novobiocin in the management of chronic hepatitis B infection. [Abstract.] Hence it would have been prima facie obvious for one of ordinary skill in the art at the time the invention was made to administer novobiocin to a mammal that is infected with hepatitis B virus. One of ordinary skill in the art at the time the invention was made would have been motivated to do so to provide treatment for hepatitis B viral infection. One of ordinary skill in the art at the time the invention was made would have had a reasonable expectoration of success for doing so because Civitico et al. demonstrates the antiviral activity of novobiocin against hepatitis B virus.

Furthermore, it would have been prima facie obvious for one of ordinary skill in the art at the time the invention was made to determine the workable or optimal treatment protocol, including dosage amount, frequency of administration, and duration of the treatment for the mammal. One of ordinary skill in the art at the time the invention was made would have been motivated to do so to treat the mammal of hepatitis B viral infection. One of ordinary skill in the art at the time the invention was made would have had a reasonable expectation of success for doing so because the determination of a workable or optimal treatment protocol is routinely practiced in the art.

⁴ Hu et al. Hsp90 is required for the activity of hepatitis B virus reverse transcriptase. Proc. Natl. Acad. Sci. USA, 1996, Vol. 93, 1060-1064.

Additionally, the Office notes that Civitico et al. does not acknowledge that the administration of novobiocin inhibits the binding of Hsp90 with its client protein or client polypeptide, hepatitis B virus reverse transcriptase. However, MPEP § 2112 [R-3] (I) provides the following:

"[T]he discovery of a previously unappreciated property of a prior art composition, or of a scientific explanation for the prior art's functioning, does not render the old composition patentably new to the discoverer." Atlas Powder Co. v. Ireco Inc., 190 F.3d 1342, 1347, 51 USPQ2d 1943, 1947 (Fed. Cir. 1999). Thus the claiming of a new use, new function or unknown property which is inherently present in the prior art does not necessarily make the claim patentable.

Thus, in accordance with the guidance provided by MPEP § 2112 [R-3] (I), Civitico et al. does not need to appreciate or acknowledge that novobiocin leads to the inhibits the binding of Hsp90 with hepatitis B virus reverse transcriptase to render the claimed invention unpatentable. In the instant, while the Office does appreciate the scientific explanation offered by Applicant, however, Applicant's discovery or appreciation of the mechanism of action in which novobiocin function is not sufficient to render Applicant's discovery patentable. In all, Applicant's appreciation or discovery of an unknown property that is inherently present in the prior art does not make Applicant's claimed invention patentable.

Furthermore, it is clearly evident from the art, Hu et al., that novobiocin does inhibit the binding of Hsp90 with hepatitis B virus reverse transcriptase to promote its antiviral activity. Hu et al. teaches that hepatitis B viruses replicates through a reverse transcription pathway. Hu et al. also teaches that the hepatitis B virus reverse transcriptase relies on Hsp90 to mediate its function. Hence, Hu et al. teaches that Hsp90 is required for the activity of hepatitis B virus reverse transcriptase, and the

Art Unit: 1648

hepatitis B virus reverse transcriptase is necessary for hepatitis B viral replication. In the instant, Civitico et al. teaches that hepatitis B viral replication is inhibited by novobiocin. Thus, novobiocin would inherently inhibit the binding of Hsp90 with hepatitis B virus reverse transcriptase, which results in the antiviral activity, inhibition of duck hepatitis B virus replication, observed by Civitico et al.

12. Claims 1 and 3-4 are rejected under 35 U.S.C. 102(b) as being anticipated by Eder et al., as evidenced by Marcu et al., as applied to claims 1 and 3 above, in view of Gormley et al.⁵

Claim 4, which depends on claim 3, which depends on claim 1, limits the coumarin or coumarin derivative to chlorobiocin or coumermycin A1.

The significance of Eder et al. and Marcu et al., as it pertains to claims 1 and 3, is provided above. As presented above, Eder et al. teaches a method of inhibiting the binding of Hsp90 with its client protein or client polypeptide in a mammal, wherein the method comprises contacting Hsp90 with novobiocin by administering about 100 mg/kg of novobiocin, for at least once per day for about 5 days to the mammal. The coumarin derivative that Eder et al. uses is novobiocin. Neither Eder et al. nor Marcu et al. teaches the use of chlorobiocin or coumermycin A1 as the coumarin derivative.

However, Gormley et al. teaches chlorobiocin, coumermycin A1 and novobiocin.

Gormley et al. teaches that chlorobiocin, coumermycin A1 and novobiocin have close structural similarity among one another, and have similar utilities. [Figure 1] Gormley et al. also teaches that chlorobiocin, coumermycin A1 and novobiocin are coumarin

⁵ Gormley et al. The interaction of coumarin antibiotics with fragments of the DNA gyrase B protein. Biochemistry, 1996, Vol. 35, 5083-5092.

Page 14

antibiotics. [Abstract] Hence, it would have been prima facie obvious for one of ordinary skill in the art at the time the invention was made to use any of these coumarin derivatives as functional alternatives for the other. One of ordinary skill in the art the time the invention was made would have been motivated to do so to inhibit the binding of Hsp90 with its client protein or client polypeptide in a mammal. One of ordinary skill in the art at the time the invention was made would have had a reasonable expectation of success for doing so because these compounds are expected to have similar properties with one another.

Response to Applicant's remarks

13. Prior to responding to Applicant's remarks, it should be noted that the rejection(s) set forth in the previous record has been withdrawn in view of the rejections submitted herein. Hence, any remarks directed at the withdrawn rejection(s) are moot.

Conclusion

- 14. No claims are allowed.
- 15. **THIS ACTION IS MADE FINAL.** Applicant is reminded of the extension of time policy as set forth in 37 CFR 1.136(a).

A shortened statutory period for reply to this final action is set to expire THREE MONTHS from the mailing date of this action. In the event a first reply is filed within TWO MONTHS of the mailing date of this final action and the advisory action is not mailed until after the end of the THREE-MONTH shortened statutory period, then the shortened statutory period will expire on the date the advisory action is mailed, and any extension fee pursuant to 37 CFR 1.136(a) will be calculated from the mailing date of

Art Unit: 1648

the advisory action. In no event, however, will the statutory period for reply expire later than SIX MONTHS from the mailing date of this final action.

16. Any inquiry concerning this communication or earlier communications from the examiner should be directed to Emily Le whose telephone number is (571) 272 0903. The examiner can normally be reached on Monday - Friday, 8 am - 5:30 pm.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Bruce R. Campell can be reached on (571) 272-0974. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic

Business Center (EBC) at 866-217-9197 (toll-free).

Art Unit 1648